

PATENT COOPERATION TREATY

From the
INTERNATIONAL SEARCHING AUTHORITY

To:
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PCT

WRITTEN OPINION OF THE
INTERNATIONAL SEARCHING AUTHORITY

(PCT Rule 43bis.1)

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| | | Date of mailing (day/month/year) 2.9 SEP 2006 |
| Applicant's or agent's file reference Y2098-10901 | | FOR FURTHER ACTION See paragraph 2 below |
| International application No. PCT/US05/47572 | International filing date (day/month/year) 30 December 2005 (30.12.2005) | Priority date (day/month/year) 30 December 2004 (30.12.2004) |
| International Patent Classification (IPC) or both national classification and IPC IPC: A61K 38/04(2006.01);C07K 5/00(2006.01),7/00(2006.01),16/00(2006.01),17/00(2006.01) USPC: 530/330 | | |
| Applicant THE TRUSTEES OF PRINCETON UNIVERSITY | | |

1. This opinion contains indications relating to the following items:

- Box No. I Basis of the opinion
- Box No. II Priority
- Box No. III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability
- Box No. IV Lack of unity of invention
- Box No. V Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement
- Box No. VI Certain documents cited
- Box No. VII Certain defects in the international application
- Box No. VIII Certain observations on the international application

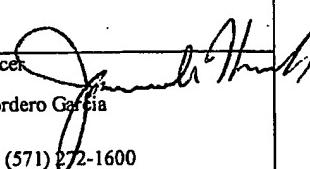
2. **FURTHER ACTION**

If a demand for international preliminary examination is made, this opinion will be considered to be a written opinion of the International Preliminary Examining Authority ("IPEA") except that this does not apply where the applicant chooses an Authority other than this one to be the IPEA and the chosen IPEA has notified the International Bureau under Rule 66.1bis(b) that written opinions of this International Searching Authority will not be so considered.

If this opinion is, as provided above, considered to be a written opinion of the IPEA, the applicant is invited to submit to the IPEA a written reply together, where appropriate, with amendments, before the expiration of 3 months from the date of mailing of Form PCT/ISA/220 or before the expiration of 22 months from the priority date, whichever expires later.

For further options, see Form PCT/ISA/220.

3. For further details, see notes to Form PCT/ISA/220.

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| Name and mailing address of the ISA/ US Mail Stop PCT, Attn: ISA/US Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450 Facsimile No. (571) 273-3201 | Date of completion of this opinion 08 September 2006 (08.09.2006) | Authorized officer  Marcela M. Cordero Garcia Telephone No. (571) 272-1600 |
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Form PCT/ISA/237 (cover sheet) (April 2005)

WRITTEN OPINION OF THE
INTERNATIONAL SEARCHING AUTHORITY

International application No.

PCT/US05/47572

Box No. I Basis of this opinion

1. With regard to the language, this opinion has been established on the basis of:
 the international application in the language in which it was filed
 a translation of the international application into _____ which is the language of a translation furnished for the purposes of international search (Rules 12.3(a) and 23.1(b)).
2. With regard to any nucleotide and/or amino acid sequence disclosed in the international application and necessary to the claimed invention, this opinion has been established on the basis of:
 - a. type of material
 a sequence listing
 table(s) related to the sequence listing
 - b. format of material
 on paper
 in electronic form
 - c. time of filing/furnishing
 contained in the international application as filed.
 filed together with the international application in electronic form.
 furnished subsequently to this Authority for the purposes of search.
3. In addition, in the case that more than one version or copy of a sequence listing and/or table(s) relating thereto has been filed or furnished, the required statements that the information in the subsequent or additional copies is identical to that in the application as filed or does not go beyond the application as filed, as appropriate, were furnished.
4. Additional comments:

WRITTEN OPINION OF THE
INTERNATIONAL SEARCHING AUTHORITY

International application No.
PCT/US05/47572

Box No. V Reasoned statement under Rule 43 bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Statement

Novelty (N) Claims 8-16 and 18-47 YES
 Claims 1-7 and 17 NO

Inventive step (IS) Claims 8-16 and 18-47 YES
 Claims 1-7 and 17 NO

Industrial applicability (IA) Claims 1-47 YES
 Claims NONE NO

2. Citations and explanations:

Claims 1-7 and 17 lack novelty under PCT Article 33(2) as being anticipated by Shyur et al. (US 7,037,696). Shyur et al. teach a compound having a polypeptide portion that includes the sequence P1-Gly-P3-Ser wherein P1 is Glu and P3 is Gly (e.g., Seq ID NO: 5, amino acids 177-180). Since the structure matches the required structure, this polypeptide reads inherently upon an inhibitor of HAUSP.

Therefore the reference anticipates the instant claims above.

Claims 8-16 and 18-47 meet the criteria set out in PCT Article 33(2)-(3), because the prior art does not teach or fairly suggest the instantly claimed products/methods.

Claims 1-47 meet the criteria set out in PCT Article 33(4), and thus have industrial applicability because the subject matter claimed can be made or used in industry.